

WHAT IS CLAIMED IS:

- 1                   1.     An isolated nucleic acid encoding a polypeptide comprising an  
2     alpha subunit of a KCNQ potassium channel, the polypeptide:
  - 3                   (i) forming, with at least one additional KCNQ alpha subunit, a  
4     KCNQ potassium channel having the characteristic of voltage-gating; and  
5                   (ii) comprising a subsequence having at least 65% amino acid  
6     sequence identity to amino acids 343 to 640 of SEQ ID NO:4.
- 1                   2.     The nucleic acid of claim 1, wherein the polypeptide specifically  
2     binds to antibodies generated against SEQ ID NO:4 or SEQ ID NO:5.
- 1                   3.     The nucleic acid of claim 1, wherein the polypeptide encodes  
2     human KCNQ5.
- 1                   4.     The nucleic acid of claim 1, wherein the nucleic acid encodes an  
2     amino acid sequence of SEQ ID NO:4 or SEQ ID NO:5.
- 1                   5.     The nucleic acid of claim 1, wherein the nucleic acid comprises a  
2     nucleotide sequence of SEQ ID NO:1, SEQ ID NO:2, or SEQ ID NO:3.
- 1                   6.     The nucleic acid of claim 1, wherein the nucleic acid is amplified  
2     by primers that selectively hybridize under stringent hybridization conditions to the same  
3     sequence as the primers selected from the group consisting of:  
4                   CCACGTCTGCACTCAGGAAGTCTCCG (SEQ ID NO:6)  
5                   CCAGCTTGGATTCTATGGACTGTACC (SEQ ID NO:7)  
6                   GAAGAGCCGAGAGAAAATAACAGCAG (SEQ ID NO:8)  
7                   GCCCTGTGGATAGCAAAGATCTTTCG (SEQ ID NO:9)  
8                   GCTGTGAGCATAAACCACTGAACCC (SEQ ID NO:10)  
9                   CCATGCGCACCATGCGGAGGATCTG (SEQ ID NO:11)  
10                  CATGAAGGATGTGGAGTCGGG (SEQ ID NO:12) and  
11                  TGGCTAAAGAACTGCTATGCCTGG (SEQ ID NO:13).
- 1                   7.     The nucleic acid of claim 1, wherein the polypeptide encoded by  
2     the nucleic acid comprises an alpha subunit of a homomeric potassium channel.

- 1                    8.        The nucleic acid of claim 1, wherein the polypeptide encoded by  
2        the nucleic acid comprises an alpha subunit of a heteromeric potassium channel.
- 1                    9.        The nucleic acid of claim 1, wherein the nucleic acid selectively  
2        hybridizes under moderately stringent hybridization conditions to a nucleotide sequence  
3        of SEQ ID NO:1, SEQ ID NO:2, or SEQ ID NO:3.
- 1                    10.      An isolated nucleic acid encoding a KCNQ polypeptide, the  
2        nucleic acid specifically hybridizing under stringent conditions to a nucleotide sequence  
3        of SEQ ID NO:1, SEQ ID NO:2, or SEQ ID NO:3.
- 1                    11.      An isolated nucleic acid that specifically hybridizes under stringent  
2        conditions to a nucleic acid encoding an amino acid sequence of SEQ ID NO:4 or SEQ  
3        ID NO:5.
- 1                    12.      A method of detecting a nucleic acid, the method comprising  
2        contacting the nucleic acid with an isolated nucleic acid of claim 1.
- 1                    13.      An isolated polypeptide comprising an alpha subunit of a KCNQ  
2        potassium channel, the polypeptide:  
3                    (i) forming, with at least one additional KCNQ alpha subunit, a  
4        KCNQ potassium channel having the characteristic of voltage-gating; and  
5                    (ii) comprising a subsequence having at least 65% amino acid  
6        sequence identity to amino acids 343 to 640 of SEQ ID NO:4.
- 1                    14.      The polypeptide of claim 13, wherein the polypeptide specifically  
2        binds to antibodies generated against SEQ ID NO:4 or SEQ ID NO:5.
- 1                    15.      The polypeptide of claim 13, wherein the polypeptide has a  
2        molecular weight of between about 95 kD to about 104 kD.
- 1                    16.      The polypeptide of claim 13, wherein the polypeptide has an amino  
2        acid sequence of human KCNQ5.
- 1                    17.      The polypeptide of claim 13, wherein the polypeptide has an amino  
2        acid sequence of SEQ ID NO:4 or SEQ ID NO:5.

1                    18.     The polypeptide of claim 13, wherein the polypeptide comprises an  
2     alpha subunit of a homomeric potassium channel.

1                    19.     The polypeptide of claim 13, wherein the polypeptide encoded by  
2     the nucleic acid comprises an alpha subunit of a heteromeric potassium channel.

1                    20.     An antibody that specifically binds to the KCNQ polypeptide of  
2     claim 13.

1                    21.     The antibody of claim 20, wherein the polypeptide to which the  
2     antibody binds has an amino acid sequence of SEQ ID NO:4 or SEQ ID NO:5.

1                    22.     An expression vector comprising the nucleic acid of claim 1.

1                    23.     A host cell transfected with the vector of claim 22.

1                    24.     A method for identifying a compound that increases or decreases  
2     ion flux through a potassium channel, the method comprising the steps of:

3                    (i) contacting the compound with a KCNQ polypeptide, the polypeptide

4                    (a) forming, with at least one additional KCNQ alpha subunit, a  
5     KCNQ potassium channel having the characteristic of voltage-gating; and

6                    (b) comprising a subsequence having at least 65% amino acid  
7     sequence identity to amino acids 343 to 640 of SEQ ID NO:4; and

8                    (ii) determining the functional effect of the compound upon the potassium  
9     channel.

1                    25.     The method of claim 24, wherein the functional effect is a physical  
2     effect.

1                    26.     The method of claim 24, wherein the functional effect is a chemical  
2     effect.

1                    27.     The method of claim 24, wherein the polypeptide is expressed in a  
2     eukaryotic host cell or cell membrane.

- 1                    28.     The method of claim 27, wherein the functional effect is  
2     determined by measuring ion flux, changes in ion concentrations, changes in current or  
3     changes in voltage.
- 1                    29.     The method of claim 24, wherein the functional effect is determined  
2     by measuring ligand binding to the channel.
- 1                    30.     The method of claim 24, wherein the polypeptide is recombinant.
- 1                    31.     The method of claim 24, wherein the potassium channel is  
2     homomeric.
- 1                    32.     The method of claim 24, wherein the potassium channel is  
2     heteromeric.
- 1                    33.     The method of claim 24, wherein the polypeptide is human KCNQ5
- 1                    34.     The method of claim 24, wherein the polypeptide has an amino acid  
2     sequence of SEQ ID NO:4 or SEQ ID NO:5.
- 1                    35.     A method of modulating ion flux through a KCNQ potassium  
2     channel, the method comprising the step of contacting the KCNQ potassium channel,  
3     wherein the channel comprises a KCNQ5 alpha subunit, with an therapeutically effective  
4     amount of a compound identified using the method of claim 24.
- 1                    36.     A method for identifying a compound that increases or decreases  
2     ion flux through a potassium channel comprising a KCNQ5 polypeptide, the method  
3     comprising the steps of:  
4                    (i) entering into a computer system an amino acid sequence of at least 50  
5     amino acids of a KCNQ5 polypeptide or at least 150 nucleotides of a nucleic acid  
6     encoding the KCNQ5 polypeptide, the KCNQ5 polypeptide comprising a subsequence  
7     having at least 65% amino acid sequence identity to amino acids 343 to 640 of SEQ ID  
8     NO:4;  
9                    (ii) generating a three-dimensional structure of the polypeptide encoded by  
10     the amino acid sequence;

11 (iii) generating a three-dimensional structure of the potassium channel  
12 comprising the KCNQ5 polypeptide;  
13 (iv) generating a three-dimensional structure of the compound; and  
14 (v) comparing the three-dimensional structures of the polypeptide and the  
15 compound to determine whether or not the compound binds to the polypeptide.

1 37. A method of detecting the presence of hKCNQ5 in human tissue,  
2 the method comprising the steps of:

3 (i) isolating a biological sample;  
4 (ii) contacting the biological sample with an hKCNQ5-specific  
5 reagent that selectively associates with hKCNQ5; and,  
6 (iii) detecting the level of hKCNQ5-specific reagent that  
7 selectively associates with the sample.

1 38. The method of claim 37, wherein the hKCNQ5-specific reagent is  
2 selected from the group consisting of: hKCNQ5-specific antibodies, hKCNQ5-specific  
3 oligonucleotide primers, and hKCNQ5-nucleic acid probes.

1 39. In a computer system, a method of screening for mutations of a  
2 human KCNQ5 gene, the method comprising the steps of:  
3 (i) entering into the computer a first nucleic acid sequence  
4 encoding a KCNQ5 polypeptide having a nucleotide sequence of SEQ ID NO:1, SEQ ID  
5 NO:2, or SEQ ID NO:3, and conservatively modified versions thereof;  
6 (ii) comparing the first nucleic acid sequence with a second nucleic  
7 acid sequence having substantial identity to the first nucleic acid sequence; and  
8 (iii) identifying nucleotide differences between the first and second  
9 nucleic acid sequences.

1 40. The method of claim 39, wherein the second nucleic acid sequence  
2 is associated with a disease state.